A large Brunner’s gland adenoma: an unusual cause of gastrointestinal bleeding
Case report and literature review

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Introduction

Although the small intestine constitutes 75% of the gastrointestinal tract, tumors are extremely rare (about 5%). Duodenum representing only 4% of the small intestine has a relatively high proportion of all the tumors as compared to the jejunum and ileum. Adenomas are the most common duodenal tumors including adenomatous polyps and Brunner adenoma 1-3. Brunner’s glands, described by the anatomist Brunner in 1688, are submucosal mucin-secreting glands. Their localization are in the duodenal bulb and progressively decrease in size and number in the distal portion 4,5. Brunner’s glands secrete an alkaline fluid composed of mucin which protect the duodenal epithelium from the acid chime of the stomach 6-9. Brunner’s gland adenoma, also known as Brunneroma or polypoid hamartoma, is a rare, benign tumor arising from the Brunner’s glands that exceptionally may evolve towards a malignant transformation 10-12. These lesions, firstly described by Curveilheir in 1835, manifest occasionally as a rare cause of duodenal obstruction or upper gastrointestinal hemorrhage, and require surgical treatment 6,8,13,14.

The present is a case of large Brunner’s gland adenoma resected by surgery with literature review. Aetiology, clinical picture, differential diagnosis and treatment of this rare tumor are discussed.

Case report

A 38-year-old female referred about six month a severe anaemia was required blood transfusions. Endoscopic examination revealed ulcer in the duodenal bulb treated by antacids and H2 blockers. When the patient was admitted to our Department of Surgical Sciences vital sign were normal but severe anaemia and vague epigastric discomfort after meals without nausea, vomiting, epigastralgia, melena or obstructive symptoms was reported. On admission she received blood transfusions, Hb 6.8 g/dl (normal 12-16 g/dl). She denied weight loss and
In 1835 Curveilheir described the first Brunner's gland adenoma situated in the proximal ileum. In our patient, the tumor was completely surrounded by duodenal mucosa. The final histological diagnosis was assessed as Brunner's gland adenoma of the duodenum. During the six month follow-up, the patient remained symptom-free with resolution of the anemia. Endoscopy did not find any residual lesion.

Discussion

In 1688 Brunner gave a precise anatomic description of the duodenal submucosal glands and used the term pancreas secondarium. In 1846 Middeldorpf correctly identified these glands as a separate entity from duodenal glands but structurally and functionally similar to glands of pylorus, and proposed be named “Brunner's glands”. Brunner's glands are mucin-secreting glands located in the deep mucosa and submucosa of duodenum, secreting an alkaline fluid composed of viscous mucus, pepsinogen and enterogastrone, whose function appears to protect the duodenal epithelium from acid of the stomach 4,6-9. They extend from pylorus distally for a variable distance, usually up to the second portion of the duodenum (Fig. 2a-b). Histological examination of the multiple biopsy obtained during endoscopy were normal (Fig. 2c). At operation, a duodenal mass was easily palpated. A longitudinal duodenotomy well distant to the pylorus removal of what appeared to be a benign tumor attached by a short peduncle with a large base to the second part (posterior and medial wall) of the duodenum. The poly was excised with its base, and the duodenotomy was closed. The surface of the tumor presented erosions and ulcers. The cut surface had a gray-red color revealing lobules. On microscopic examination the tumor was composed of hyperplasia of Brunner's glands. The hyperplastic glands formed lobules and ducts embedded in a fibrous stroma, adipose tissue, and aggregated lymphoid. No signs of malignancy were found and the tumor was completely surrounded by duodenal mucosa. The final histological diagnosis was assessed as Brunner's gland adenoma of the duodenum. During the six month follow-up, the patient remained symptom-free with resolution of the anemia. Endoscopy did not find any residual lesion.

any recent intake of non-steroidal anti-inflammatory drugs. The physical examination had no remarkable finding. The abdomen was soft without palpating pain and jumping pain. An ultrasonography was carried out showing the presence of a endoluminal mass in the first and second portion of the duodenum measuring 5.8 x 4.8 cm (Fig. 1). On upper endoscopy demonstrated normal esophagus and stomach. A large poly with a short peduncle completely occupying the duodenal lumen was seen, protruding between the bulb and the second part of the duodenum (Fig. 2a-b). Histological examination of the multiple biopsy obtained during endoscopy were normal (Fig. 2c). At operation, a duodenal mass was easily palpated. A longitudinal duodenotomy well distant to the pylorus removal of what appeared to be a benign tumor attached by a short peduncle with a large base to the second part (posterior and medial wall) of the duodenum. The poly was excised with its base, and the duodenotomy was closed. The surface of the tumor presented erosions and ulcers. The cut surface had a gray-red color revealing lobules. On microscopic examination the tumor was composed of hyperplasia of Brunner's glands. The hyperplastic glands formed lobules and ducts embedded in a fibrous stroma, adipose tissue, and aggregated lymphoid. No signs of malignancy were found and the tumor was completely surrounded by duodenal mucosa. The final histological diagnosis was assessed as Brunner's gland adenoma of the duodenum. During the six month follow-up, the patient remained symptom-free with resolution of the anemia. Endoscopy did not find any residual lesion.

Autopsies and Brunner's gland adenoma comprises 10.6% of these lesions 17-20. Size of the tumor is important in differentiating adenoma from Brunner gland hyperplasia. The size less than 1 cm is referred to as Brunner gland hyperplasia 3,5,14,16.

Aetiology and pathogenesis of Brunner's gland adenoma still remains elusive, but associations with peptic ulcer disease, chronic pancreatitis and chronic renal insufficiency have been described 5,6. Brunner's gland hyperplasia is believed to act as a mechanism which allow to protect the duodenal mucosa 14,17. Given that Brunner's glands have an anti-acid function, it has been postulated that gastric hyperacidity could stimulate these structures to undergo hyperplasia, but only 45% of patients had hyperacidity and 20% had hypoacidity 6,20-22. Franzin et al have reported an association between Brunner's gland adenoma and gastric hyperacidity in patients with chronic gastric erosions and duodenal ulcers. Nevertheless Spellberg et al have not found regression of the lesion with acid secretion inhibitors. Other mechanism suggested is inflammatory origin due to the presence of a dense inflammatory cell infiltration, but given that lymphocytes are usually present in the normal submucosa, the presence of inflammatory cell in the Brunner's gland adenoma is not sufficient to substan this inflammatory theory 14,21,25. Finally, it has suggested that Helicobacter pylori infection may play a role in the pathogenesis of Brunner's gland adenoma, but its role remains unclear 26,27. In our opinion, on the whole of lobules of Brunner's glands separated by fibrovascular septa embedded in a fibrous stroma with adipose tissue and lymphocyte and monocyte infiltration, suggests to identify this lesion as a hamartoma or nodular hyperplasia rather than a real neoplasm. Although the malignant degeneration is rare, two cases have been reported, including one patient with microcarcinoid tumors 8,28. Fujimaki et al reported one patient with a focus atypical gland and p-53 positive cells detected sporadically and conclude that the apical glands probably represented a neoplastic lesion. The most common location for the Brunner's gland adenoma is the posterior wall of the duodenal bulb (57%) 29. In the majority of cases, this lesion present into a polypoid mass usually pedunculated (88%) 20. Brunner's gland adenoma was rarely found extending to the proximal jejunum 8,21. Laarman et al reported a Brunner's gland adenoma located in the proximal jejunum ; Hanken and Forouhar described a patient with a Brunner's gland adenoma situated in the proximal ileum. In our patient, Brunner's gland adenoma are located in the second part of the duodenum (posterior and medial wall).

Feyrter classified three types of Brunner's gland hyperplasia: diffuse nodular hyperplasia (Type I) confined to the mucosa with multiple sessile projections occupying most of the duodenum; circumscribed nodular hyperplasia (Type II) the most common type mainly present in the duodenal bulb, usually less than 1 cm in size; adenomatous hyper-
plasia (Type III), also known as Brunner’s gland hamartoma or adenoma or Brunneroma, which generally present as a single polypoid lesion, sessile or pedunculated. Hyperplasia refers to multiple lesions (usually less than 1 cm) and adenoma refers to a lesion larger than 1 cm. Brunner’s gland adenoma has a tendency to be predominant in the fifth or sixth decade of life with equal gender distribution. Symptoms of the patients with Brunner’s gland adenoma can be inserted into two categories: symptomatic tumors that can further be divided into haemorrhagic and obstructive tumors; and asymptomatic tumors that are only found incidentally. The most common clinical manifestations of symptomatic group are gastrointestinal hemorrhage, due to ulceration or erosion of the tumor. Obstructive tumors causing epigastric pain, discomfort, vomiting or weight loss. There are also reports about patients who complained of diarrhoea owing to disturbances of duodenal motility. Levine et al studied the characteristics of a group of 27 patients with Brunner’s gland adenoma and described three types of clinical picture. An asymptomatic group comprising 11% in whom the Brunner’s gland adenoma was an incidental finding. The second group (40-50%) consisted of patients with upper gastrointestinal bleeding. These patients with bleeding complications present with melaena, fatigue, malaise and anaemia. Although blood loss is usually occult, massive and even fatal haemorrhage has been described. The last group comprised patients with prolonged obstructive upper gastrointestinal symptoms. These obstructive symptoms is present in about 50% of patients, giving rise to epigastric pain, nausea, vomiting and postprandial discomfort. Duodenal intussusception, pancreatitis and diarrhoea are rare presentations. Our patient presented initially with severe anaemia and later on vague epigastric discomfort but no obstructive symptoms although the lesion was too large. Preoperative histological diagnosis of Brunner’s gland adenoma is not always easy at present. Diagnostic radiographic studies include especially ultrasonography, barium contrast studies and endoscopy. Large adenomas may be detected by ultrasonography. Upper gastrointestinal barium studies may reveal multiple small filling defect (Swiss cheese appearance) in Brunner’s gland hyperplasia or, in the case of an adenoma, a smooth polypoid defect may be seen. Hypotonic duodenography may have a role for delineating the surface of the lesion. In X-rays barium examination, the findings are often non-specific because there is usually a sessile or pedunculated polypoid-filling defect in the duodenal bulb. Upper gastrointestinal endoscopy with biopsies is the diagnostic method of choice. She has an additional function in diagnosing and treating Brunner’s gland adenoma, since than is possible to verify the histological diagnosis and remove the tumor at the same time. Endoscopic pinching biopsies is usually negative since the biopsy forceps are unable to reach the tumoral tissue localized completely in the submucosa layer. Computed tomography is useful only to confirm the absence of extraluminal extension of Brunner’s gland adenoma. In our case, first the
tumor was detected by ultrasonography and then confirmed by endoscopic examination; biopsies, however, showed normal tissue and the final diagnosis was made with pathological examination. Brunner’s adenoma can mimic several other lesions; the differential diagnosis includes benign tumours of the small intestine such as adenoma of the Islet cells, polypoid adenoma, leiomyoma, lipoma, angioma, aberrant pancreatic tissue, transudated mucosa, duodenal duplication cyst and malignant tumours as adenocarcinoma, lymphoma, carcinoids tumour and leiomyosarcoma.

Brunner’s gland adenomas are considered a benign disorder; malignant degeneration has only been described in literature. It is still controversial whether Brunner’s gland adenoma can be removed endoscopically or with transduodenal exploration and excision of the polyp by laparoscopic or open surgery. Endoscopic polypectomy represents the ideal approach when the tumour is pedunculated and still relatively small. However, the success depends by an experienced surgeon. Open surgical excision with transduodenal exploration, excision of the polyp and suture of duodenotomy, is reserved for cases where endoscopic treatment has failed or, like our, when the tumour is too large.

In our case, because of site and size of the lesion (5.8 cm x 4.8 cm) with a very short peduncle occupying entirely the duodenal bulb, surgical removal was effective for the treatment of Brunner’s gland adenoma without complications and completely resolution of the anemia. Moreover endoscopy did not find any residual lesion or recurrence during six month follow-up.

Conclusion

Brunner’s gland adenoma is a rare benign tumour of the duodenum and itself is a very rare and insidious cause of upper gastrointestinal bleed. Patients may also present with symptoms of obstruction. Endoscopic polypectomy is the first choice when the tumour is small or pedunculated. Surgical excision is reserved for cases where endoscopic treatment has failed or when tumour is too large.

Riassunto

L’adenoma delle ghiandole di Brunner è un raro tumore benigno del duodeno. La sua più frequente localizzazione è a livello della parete posteriore tra prima e seconde porzione duodenale. Generalmente piccolo ed asintomatico quando viene scoperto accidentalmente in corso di esame endoscopico, a volte può essere di grandi dimensioni e causare sintomi emorragici od ostruttivi. Gli Autori riportano il caso clinico di un adenoma di Brunner di grandi dimensioni in una donna di 38 anni presentatosi con un quadro di anemia severa ma senza sintomi ostruttivi e scoperto mediante indagine ecografica. Il tumore è stato asportato chirurgicamente e la paziente a sei mesi di follow-up è risultata in buona salute, libera da malattia e con completa risoluzione del quadro anemico. È stata inoltre eseguita una revisione della più recente letteratura.

References


